Claims

1. A compound of formula (I) or a pharmaceutically acceptable salt thereof:

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$$(R_1)_p$$
 Q (I)

wherein:

 R_1 is halogen, cyano, C_{1-6} alkyl, C_{3-7} cycloalkyl, C_{3-7} cycloalkyloxy, C_{1-6} alkoxy, C_{1-6} alkylthio, hydroxy, amino, mono- or di- C_{1-6} alkylamino, an N-linked 4 to 7 membered heterocyclic group, nitro, halo C_{1-6} alkyl, halo C_{1-6} alkoxy, aryl, -COOR₃, -COR₄ (wherein R_3 and R_4 , are independently hydrogen or C_{1-6} alkyl) or -COR₅ (wherein R_5 is amino, mono- or di- C_{1-6} alkylamino or an N-linked 4 to 7 membered heterocyclic group); p is 0, 1 or 2 or 3;

Q is a 6- membered aromatic group or a 6-membered heteroaromatic group;

A is -(CH₂-CH₂)-, -(CH=CH)-, or a group -(CHR₇)- wherein R₇ is hydrogen, halogen, hydroxy, cyano, nitro, C₁₋₆alkyl, C₃₋₇cycloalkyl, C₃₋₇cycloalkyloxy, haloC₁₋₆alkyl, C₁₋₆alkoxy, haloC₁₋₆alkoxy or C₁-6alkylthio;

 R_2 is hydrogen, halogen, hydroxy, cyano, nitro, C_{1-6} alkyl, C_{1-6} alkanoyl, C_{3-7} cycloalkyl, C_{3-7} cycloalkyl, C_{1-6} alkoxy, halo C_{1-6} alkoxy, halo C_{1-6} alkylthio, amino, mono- or di- C_{1-6} alkylamino or an N-linked 4 to 7 membered heterocyclic group;

X is oxygen, sulfur, -CH₂- or NR₈ wherein R₈ is hydrogen or C₁₋₆alkyl;

Y is a single bond, -CH₂-, -(CH₂)₂- or -CH=CH-; and

Z is an optionally substituted N-linked heterocyclic group or a C-linked 4 to 7 membered heterocyclic group containing at least one nitrogen, or Z is $-NR_9R_{10}$ where R_9 and R_{10} are independently hydrogen or C_{1-6} alkyl.

- 2. A compound as claimed in claim 1, wherein when R₇ is hydrogen.
- 3. A compound as claimed in claim 1 or claim 2, wherein A is CH₂-.

4. A compound as claimed in any of claims 1-3, wherein Q is phenyl.

5. A compound as claimed in any of claims 1-4, wherein p is 1, 2 or 3, and R_1 is/are halogen (particularly chloro or fluoro), C_{1-6} alkyl (particularly methyl) or CF_3 .

6. A compound as claimed in any of claims 1-5, wherein when R_1 is attached at the position marked below with an asterisk, R_1 is fluoro:

7. A compound as claimed in any of claims 1-6, wherein when Q is phenyl and p is 1, R_1 is attached at the position marked below with an asterisk:

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8. A compound as claimed in any of claims 1-7, wherein when Q is phenyl and p is 2 or 3, R₁ is attached at two or more of the positions marked below with arrows:

- 9. A compound as claimed in any of claims 1-8, wherein R_2 is C_{1-6} alkoxy, 15 particularly methoxy.
 - 10. A compound as claimed in any of claims 1-9, wherein X is oxygen.

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- 11. A compound as claimed in any of claims 1-10, wherein Y is -CH₂-.
- 12. A compound as claimed in any of claims 1-11, wherein Z is an optionally substituted N-linked 4 to 7 membered heterocycle, in particular optionally substituted piperidyl.
- 25 13. A compound as claimed in claim 1 having the formula (la):

$$(R_1)_p$$
 (Ia)

wherein R_1 , p, R_2 , X, Y, Z, are as defined in any of claims 1-12 and A_1 is -CH₂- or -HC(Me)-.

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14. A compound as claimed in claim 1, which is

2-[4-Methoxy-3-(2-piperidin-1-yl-ethoxy)phenyl]-2,3-dihydroisoindol-1-one 6-Fluoro-2-[4-methoxy-3-(2-piperidin-1-yl-ethoxy)phenyl]-2,3-dihydroisoindol-1-one

- 7-Bromo-2-{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy]phenyl}-2,3-dihydroisoindol-1-one hydrochloride
 - 7-Chloro-2-{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy]phenyl}-3-methyl-2,3-dihydroisoindol-1-one
 - 2-{4-Methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy]phenyl}-7-trifluoromethyl-2,3-
- 15 dihydroisoindol-1-one
 - 5,7-Dichloro-2-{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy]phenyl}-2,3-dihydroisoindol-1-one
 - 7-Chloro-2-[4-methoxy-3-(2-piperidin-1-yl-ethoxy)phenyl]-2,3-dihydroisoindol-1-one 6-Chloro-2-[4-methoxy-3-(2-piperidin-1-yl-ethoxy)phenyl]-2,3-dihydroisoindol-1-one
- 20 hydrochloride

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- 5-Chloro-2-[4-methoxy-3-(2-piperidin-1-yl-ethoxy)phenyl]-2,3-dihydroisoindol-1-one 5,7-Dichloro-2-{4-methoxy-3-[2-(*cis*-2,6-dimethyl-piperidin-1-yl)-ethoxy]phenyl}-2,3-dihydroisoindol-1-one
- 7-Chloro-2-{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy]phenyl}-2,3-dihydroisoindol-1-one
- 6-Chloro-2-{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy]phenyl}-2,3-dihydroisoindol-1-one
- $5- Chloro-2-\{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy] phenyl\}-2, \\ 3-dihydroisoindol-1-one$
- 30 7-Methyl-2-{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy]phenyl}-2,3-dihydroisoindol-1-one
 - $6,7-Difluoro-2-\{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy] phenyl\}-2,3-dihydroisoindol-1-one$
 - 5,6-Dichloro-2-{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy]phenyl}-2,3-dihydroisoindol-1-one
 - 7-Fluoro-2-{4-methoxy-3-[2-(piperidin-1-yl)-ethoxy]phenyl}-2,3-dihydroisoindol-1-one 4-Fluoro-2-{4-methoxy-3-[2-(piperidin-1-yl)-ethoxy]phenyl}-2,3-dihydroisoindol-1-one

5,7-Dimethyl-2-{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy]phenyl}-2,3-dihydroisoindol-1-one

- $6,7- Dichloro-2-\{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy] phenyl\}-2,3-dihydroisoindol-1-one$
- 5 5-Fluoro-2-{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy]phenyl}-7-trifluoromethyl-2,3-dihydroisoindol-1-one
 - $7-Chloro-4, 5-difluoro-2-\{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy] phenyl\}-2, 3-dihydroisoindol-1-one$
 - 4-Fluoro-2-{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy]phenyl}- 7-trifluoromethyl-2,3-
- dihydroisoindol-1-one

 4-Fluoro-2-{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy]phenyl}- 7-trifluoromethyl-2,3-dihydroisoindol-1-one
 - 4-Fluoro-2-{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy]phenyl}- 7-trifluoromethyl-2,3-dihydroisoindol-1-one
- 4-Fluoro-2-{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy]phenyl}- 7-trifluoromethyl-2,3-dihydroisoindol-1-one
 - $5,7\text{-}Dichloro-2-\{4\text{-}methoxy-3-[2-(4,4\text{-}dimethyl\text{-}piperidin-1-yl)-ethoxy]} phenyl\}-2,3-dihydroisoindol-1-one$
 - 5,7-Dichloro-2-{4-methoxy-3-[2-(azepan-1-yl)-ethoxy]phenyl}-2,3-dihydroisoindol-1-one
- 5,7-Dichloro-2-{4-methoxy-3-[2-(2-methyl-piperidin-1-yl)-ethoxy]phenyl}-2,3-dihydroisoindol-1-one
 - $6-\{4-Methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy]-phenyl\}-2-methyl-4-trifluoromethyl-6, 7-dihydro-pyrrolo[3,4-b]pyridin-5-one \\$
 - 5,7-Dichloro-4-fluoro-2-{4-methoxy-3-[2-(4-methyl-piperidin-1-yl)-ethoxy]phenyl}-2,3-dihydroisoindol-1-one
- or a pharmaceutically acceptable salt thereof.
 - 15. A process for the preparation of a compound as claimed in any of claims 1-14 or a pharmaceutically acceptable salt thereof, which process comprises:
- 30 (a) reacting a compound of formula (II):

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$$(R_1)_p$$
 (II)

wherein R₁, R₂, p, A, X, and Y are as defined for formula (I), and L is a leaving group, with a compound of formula (III):

Z-H (III)

wherein Z is as defined for formula (I); or

(b) reacting a compound of formula (IV):

$$(R_1)_p$$
 ORX (IV)

wherein Rx is alkyl and LG is a suitable leaving group, with a compound of formula (V) or a corresponding salt:

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or

(c) reacting a compound of formula (VI):

$$(R_1)_p$$
 (VI)

with a compound of formula (V) in the presence of AlMe₃ or a similar oxophilic reagent followed by treatment of the resulting amide under dehydrating conditions, e.g. with PPh₃ and dialkylazadicarboxylate;

and thereafter, for either process (a), process (b) or process (c), optionally followed by:

- removing any protecting groups; and/or
- converting a compound of formula (I) into another compound of formula (I); and/or
- forming a pharmaceutically acceptable salt.

16. A pharmaceutical composition comprising a compound as defined in any of claims 1-14 or a pharmaceutically acceptable salt thereof, and a pharmaceutically acceptable carrier or excipient.

- 5 17. A process for preparing a pharmaceutical composition as defined in claim 16, the process comprising mixing a compound as defined in any of claims 1-14 or a pharmaceutically acceptable salt thereof and a pharmaceutically acceptable carrier or excipient.
- 10 18. A compound as defined in any of claims 1-14 or a pharmaceutically acceptable salt thereof for use as a therapeutic substance.

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- 19. A compound as defined in any of claims 1-14 for use in the treatment of a CNS disorder such as depression or anxiety.
- 20. A method of treatment of a CNS disorder in mammals including humans, which comprises administering to the sufferer a therapeutically safe and effective amount of a compound as claimed in any of claims 1-14 or a pharmaceutically acceptable salt thereof.
- 20 21. A method as claimed in claim 20, wherein the CNS disorder is depression or anxiety.
- 22. Use of a compound as claimed in any of claims 1-14 or a pharmaceutically acceptable salt thereof in the manufacture of a medicament for use in the treatment of a CNS disorder.
 - 23. The use as claimed in claim 22, wherein the CNS disorder is depression or anxiety.